Atty Dkt. No.: GRUE003

USSN: 09/269,874

II. REMARKS

Formal Matters

The specification is amended to refer to sequence identifiers. Support for the amendments is found in the specification and figures at, inter alia, Example 1, section A, page 15. Accordingly, no new matter is introduced by the amendments to the specification.

Attached hereto is a marked-up version of the changes made to the specification by the current amendment. The attached is captioned "VERSION WITH MARKINGS TO SHOW CHANGES MADE."

Applicants respectfully entry of the above-noted amendments to the specification.

Objections to the specification

This communication is responsive to the Office Action which issued on September 13, 2001. The Office Action stated that the instant specification does not comply with the requirements of 37 C.F.R.§1.821-1.825. Applicants respectfully request the above-noted amendments to the specification.

Sequence Listing

A new Sequence Listing is provided herewith, which is in compliance with the requirements of 37 C.F.R.§1.821-1.825. No new matter is added.

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III. CONCLUSION

If the Examiner finds that a telephone conference would expedite the prosecution of this application, the Examiner is invited to telephone the undersigned at the number provided.

The Commissioner is hereby authorized to charge any underpayment of fees associated with this communication, including any necessary fees for extensions of time, or credit any overpayment to Deposit Account No. 50-0815, order number GRUE-003.

Respectfully submitted, BOZICEVIC, FIELD & FRANCIS LLP

Date: Oct. 12, 2001

Paula A. Borden

Registration No. 42,344

BOZICEVIC, FIELD & FRANCIS LLP 200 Middlefield Road, Suite 200 Menlo Park, CA 94025

Telephone: (650) 327-3400 Facsimile: (650) 327-3231

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VERSION WITH MARKINGS TO SHOW CHANGES MADE

Please enter the following amendments to the specification.

Please amend page 14, line 2, as follows:

[Total sequencing of gp190^s] <u>Figure 3C provides the native nucleotide sequence (gp190ⁿ; SEQ ID NO:1) encoding gp190; the synthetic nucleotide sequence (gp190^s; SEQ ID NO:2) encoding gp190; and the amino acid (SEQ ID NO:3) of gp190.</u>

Please amend page 14, line 3, as follows:

N- and C-termini of gp190^{S1} and gp190^{S2} variants [variant]

Please amend page 16, line 16, as follows:

D. N- and C-termini of gp190^{S1} and gp190^{S2} variants [variant] (see Fig. 3D)

Please amend the paragraph beginning on page 16, line 22, as follows:

The N-terminal and C-terminal sequences of gp120^{S1} are provided as SEQ ID NO:4 and SEQ ID NO:5, respectively. Figure 3D depicts nucleotides 1-17 and 4863-4894 of SEQ ID NO:4 and amino acids 1-3 and 1619-1621 of SEQ ID NO:5. The N-terminal sequence of gp120^{S2}, beginning with the BamHI cleavage site, indicates the transition at amino-acid 20, from which it can be assumed that after splitting of the signal peptide it defines the N-terminus. At the C-terminus the sequence encoded ended at amino-acid 1621. The stop-codon followed the ClaI cleavage site. The nucleotide and amino acid sequences of gp120^{S2} are provided as SEQ ID NO:6 and SEQ ID NO:7, respectively. Figure 3D depicts nucleotides 1-17 and 4806-4838 of SEQ ID NO:6 and amino acids 1-3 and 1600-1602 of SEQ ID NO:7.

Please amend the paragraph beginning on page 17, line 3, as follows:

The gp120^{S2} sequence was inserted via the BamHI and ClaI cleavage sites into pDS56RBSII, by means of which 6 histidine as well as some amino-acids originating in the vector were fused to the N-terminus. This produces the following N-terminal sequence on the reading-frame:

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 $\label{eq:metargGlySer} MetArgGlySer(His)_6GlySer~\underbrace{(SEQ~ID~NO:8)}_{.}.~~ Through the promoter~P_{N25lac0-1}~ the transcription comes~\\ under lacR/O/IPTG~control.$

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NOTICE TO COMPLY WITH REQUIREMENTS FOR PATENT APPLICATIONS CONTAINING NUCLEOTIDE SEQUENCE AND/OR AMINO ACID SEQUENCE DISCLOSURES

The nucleotide and/or amino acid sequence disclosure contained in this application does not comply with the requirements for such a disclosure as set forth in 37 C.F.R. 1.821 - 1.825 for the following reason(s):



- 1. This application clearly fails to comply with the requirements of 37 C.F.R. 1.821-1.825. Applicant's attention is directed to these regulations.
- 2. This application does not contain, as a separate part of the disclosure on paper copy, a "Sequence Listing" as required by 37 C.F.R. 1.821(c).
- 3. A copy of the "Sequence Listing" in computer readable form has not been submitted as required by 37 C.F.R. 1.821(e).
- 4. A copy of the "Sequence Listing" in computer readable form has been submitted. However, the content of the computer readable form does not comply with the requirements of 37 C.F.R. 1.822 and/or 1.823, as indicated on the attached copy of the marked -up "Raw Sequence Listing."
- 5. The computer readable form that has been filed with this application has been found to be damaged and/or unreadable as indicated on the attached CRF Diskette Problem Report. A Substitute computer readable form must be submitted as required by 37 C.F.R. 1.825(d).
- 6. The paper copy of the "Sequence Listing" is not the same as the computer readable form of the "Sequence Listing" as required by 37 C.F.R. 1.821(e).
- 7. Other: All disclosed sequences are not listed as required by 37 CFR § 1.821(c), because the sequences disclosed at least on page 17, line 6, and in Fig. 3D do not appear in either the CRF or the paper copy of the "Sequence Listing". Also, "SEQ ID NO:" identifiers must be entered as required by 37 CFR § 1.821(d).

Applicant Must Provide:

An initial or substitute computer readable form (CRF) copy of the "Sequence Listing".

An initial or substitute paper copy of the "Sequence Listing", as well as an amendment directing its entry into the specification.

A statement that the content of the paper and computer readable copies are the same and, where applicable, include no new matter, as required by 37 C.F.R. 1.821(e) or 1.821(f) or 1.821(g) or 1.825(b) or 1.825(d).

Amendments to the specification directing entry of "SEQ ID NO:" identifiers into the specification.

For questions regarding compliance to these requirements, please contact:

For Rules Interpretation, call (703) 308-4216

For CRF Submission Help, call (703) 308-4212

PatentIn Software Program Support (SIRA)

PLEASE RETURN A COPY OF THIS NOTICE WITH YOUR RESPONSE





SEQUENCE LISTING

OCT 2 2 2001

TECH CENTER 1600/2900

1620

<120> Recombinant Process for Preparing a Complete Malaria Antigen, GP190/MSP1

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Pro Gln Leu Phe Asp Leu Thr Asn His Met Leu Thr Leu Cys Asp Asn Ile His Gly Phe Lys Tyr Leu Ile Asp Gly Tyr Glu Glu Ile Asn Glu Leu Leu Tyr Lys Leu Asn Phe Tyr Phe Asp Leu Leu Arg Ala Lys Leu Asn Asp Val Cys Ala Asn Asp Tyr Cys Gln Ile Pro Phe Asn Leu Lys Ile Arg Ala Asn Glu Leu Asp Val Leu Lys Lys Leu Val Phe Gly Tyr Arg Lys Pro Leu Asp Asn Ile Lys Asp Asn Val Gly Lys Met Glu Asp Tyr Ile Lys Lys Asn Lys Lys Thr Ile Glu Asn Ile Asn Glu Leu Ile Glu Glu Ser Lys Lys Thr Ile Asp Lys Asn Lys Asn Ala Thr Lys Glu Glu Glu Lys Lys Leu Tyr Gln Ala Gln Tyr Asp Leu Ser Ile Tyr Asn Lys Gln Leu Glu Glu Ala His Asn Leu Ile Ser Val Leu Glu Lys Arg Ile Asp Thr Leu Lys Lys Asn Glu Asn Ile Lys Glu Leu Leu Asp Lys Ile Asn Glu Ile Lys Asn Pro Pro Pro Ala Asn Ser Gly Asn Thr Pro Asn Thr Leu Leu Asp Lys Asn Lys Ile Glu Glu His Glu Lys Glu Ile Lys Glu Ile Ala Lys Thr Ile Lys Phe Asn Ile Asp Ser Leu Phe Thr Asp Pro Leu Glu Leu Glu Tyr Tyr Leu Arg Glu Lys Asn Lys Asn Ile Asp Ile Ser Ala Lys Val Glu Thr Lys Glu Ser Thr Glu Pro Asn Glu Tyr Pro Asn Gly Val Thr Tyr Pro Leu Ser Tyr Asn Asp Ile Asn Asn Ala Leu Asn Glu Leu Asn Ser Phe Gly Asp Leu Ile Asn Pro Phe Asp Tyr Thr Lys Glu Pro Ser Lys Asn Ile Tyr Thr Asp Asn Glu Arg Lys Lys Phe Ile Asn Glu Ile Lys Glu Lys Ile Lys Ile Glu Lys Lys Lys Ile Glu Ser Asp Lys Lys Ser Tyr Glu Asp Arg Ser Lys Ser Leu Asn Asp Ile Thr Lys Glu Tyr Glu Lys Leu Leu Asn Glu Ile Tyr Asp Ser Lys Phe Asn Asn Asn Ile Asp Leu Thr Asn Phe Glu Lys Met Met Gly Lys Arg Tyr Ser Tyr Lys Val Glu Lys Leu Thr His His Asn Thr Phe Ala Ser Tyr Glu Asn Ser Lys His Asn Leu Glu Lys Leu Thr Lys Ala Leu Lys Tyr Met Glu Asp Tyr Ser Leu Arg Asn Ile Val Val Glu Lys Glu Leu Lys Tyr Tyr Lys Asn Leu Ile Ser Lys Ile Glu Asn Glu Ile Glu Thr Leu Val Glu Asn Ile Lys Lys Asp Glu Glu Gln Leu Phe Glu Lys Lys Ile Thr Lys Asp Glu Asn Lys Pro Asp Glu Lys Ile

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Asn Lys Gln Glu Pro Tyr Tyr Leu Ile Val Leu Lys Lys Glu Ile Asp
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Lys Leu Lys Val Phe Met Pro Lys Val Glu Ser Leu Ile Asn Glu Glu
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Lys Lys Asn Ile Lys Thr Glu Gly Gln Ser Asp Asn Ser Glu Pro Ser
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Ala Gln Val Pro Thr Pro Pro Ala Pro Val Asn Asn Lys Thr Glu Asn
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Ile Pro Val Met Tyr Ser Met Phe Asp Ser Leu Asn Asn Ser Leu Ser
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Gln Leu Phe Met Glu Ile Tyr Glu Lys Glu Met Val Cys Asn Leu Tyr
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                                           860
Lys Leu Lys Asp Asn Asp Lys Ile Lys Asn Leu Leu Glu Glu Ala Lys
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Gln Lys Ser Ser Glu Asn Phe Tyr Glu Lys Ile Leu Lys Asp Ser Asp
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Asp Ile Asn Lys Leu Lys Lys Thr Leu Gln Leu Ser Phe Asp Leu Tyr
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                                               1005
Asn Lys Tyr Lys Leu Lys Leu Glu Arg Leu Phe Asp Lys Lys Thr
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                       1015
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Val Gly Lys Tyr Lys Met Gln Ile Lys Lys Leu Thr Leu Leu Lys Glu
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Gln Leu Glu Ser Lys Leu Asn Ser Leu Asn Asn Pro Lys His Val Leu
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1505 1510 1515 Leu Gln Gly Met Leu Asn Ile Ser Gln His Gln Cys Val Lys Lys Gln 1525 1530 Cys Pro Gln Asn Ser Gly Cys Phe Arg His Leu Asp Glu Arg Glu Glu 1545 1550 Cys Lys Cys Leu Leu Asn Tyr Lys Gln Glu Gly Asp Lys Cys Val Glu 1555 1560 1565 Asn Pro Asn Pro Thr Cys Asn Glu Asn Asn Gly Gly Cys Asp Ala Asp 1570 1575 1580 Ala Lys Cys Thr Glu Glu Asp Ser Gly Ser Asn Gly Lys Lys Ile Thr 1590 1595 Cys Glu Cys Thr Lys Pro Asp Ser Tyr Pro Leu Phe Asp Gly Ile Phe 1605 1610 Cys Ser Ser Ser Asn Phe Leu Gly Ile Ser Phe Leu Leu Ile Leu Met 1620 1625 Leu Ile Leu Tyr Ser Phe Ile 1635

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Glu Lys Met Val Leu Asn Glu Gly Thr Ser Gly Thr Ala Val Thr Thr
Ser Thr Pro Gly Ser Lys Gly Ser Val Ala Ser Gly Gly Ser Gly Gly
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Ser Val Ala Ser Gly Gly Ser Val Ala Ser Gly Gly Ser Val Ala Ser
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Gly Gly Ser Val Ala Ser Gly Gly Ser Gly Asn Ser Arg Arg Thr Asn
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Pro Ser Asp Asn Ser Ser Asp Ser Asp Ala Lys Ser Tyr Ala Asp Leu
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                            120
Lys His Arg Val Arg Asn Tyr Leu Leu Thr Ile Lys Glu Leu Lys Tyr
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Pro Gln Leu Phe Asp Leu Thr Asn His Met Leu Thr Leu Cys Asp Asn
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Ile His Gly Phe Lys Tyr Leu Ile Asp Gly Tyr Glu Glu Ile Asn Glu
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Leu Leu Tyr Lys Leu Asn Phe Tyr Phe Asp Leu Leu Arg Ala Lys Leu
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Arg Lys Pro Leu Asp Asn Ile Lys Asp Asn Val Gly Lys Met Glu Asp
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Tyr Ile Lys Lys Asn Lys Lys Thr Ile Glu Asn Ile Asn Glu Leu Ile
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Arg Ile Asp Thr Leu Lys Lys Asn Glu Asn Ile Lys Glu Leu Leu Asp
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